

CLAIM AMENDMENTS

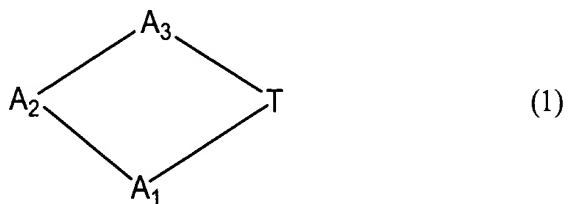
Listing of Claims:

Claims 1-23 (canceled)

Claims 24-33 (canceled)

Claim 34 (not entered)

Claim 35 (new): A macrocyclic compound of the formula (1):

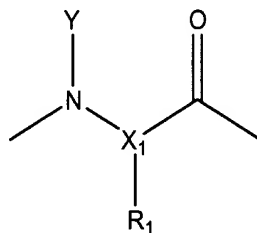


and its pharmaceutically acceptable salts,

wherein

Fragment A₁ is:

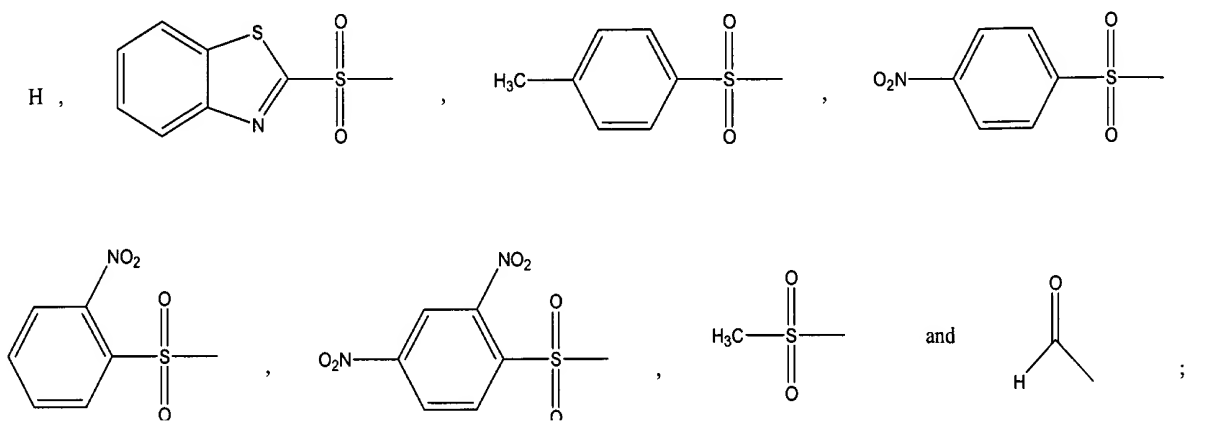
(1-i)



wherein

Y is selected from the group consisting of

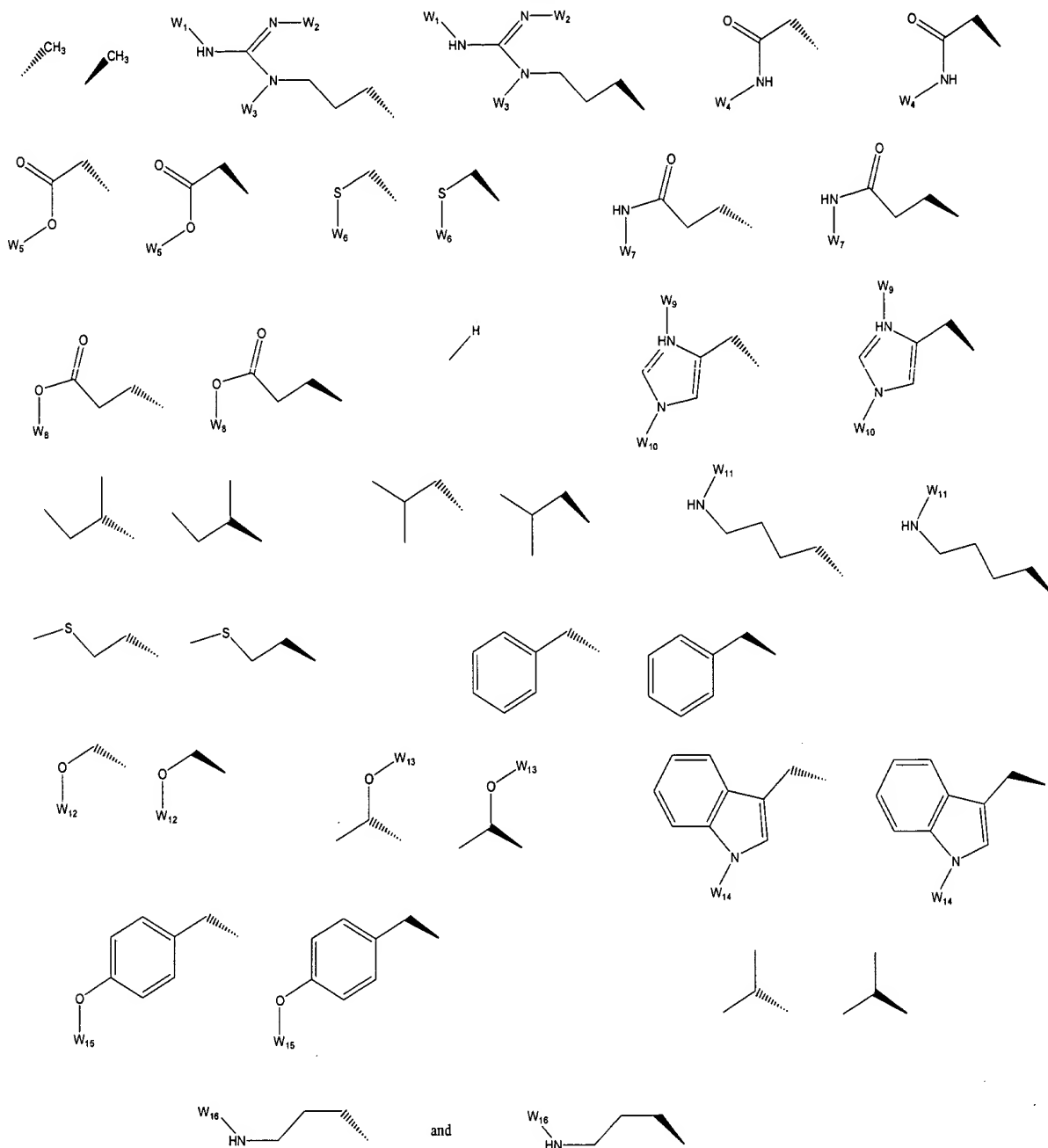
{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



X_1 is $-\text{CH}-$, $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$;

when X_1 is $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$, R_1 is absent;

when X_1 is $-\text{CH}-$, R_1 is a radical independently selected from the group consisting of

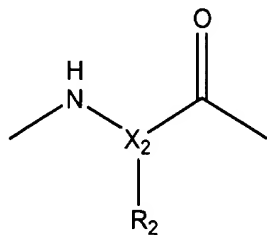


Fragment A₂ is:

(2-i) *D*-proline, *L*-proline, *D*-4-hydroxyproline, *L*-4-hydroxyproline, *D*-4-tert-butoxyproline, *L*-4-tert-butoxyproline; or

(2-ii)

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

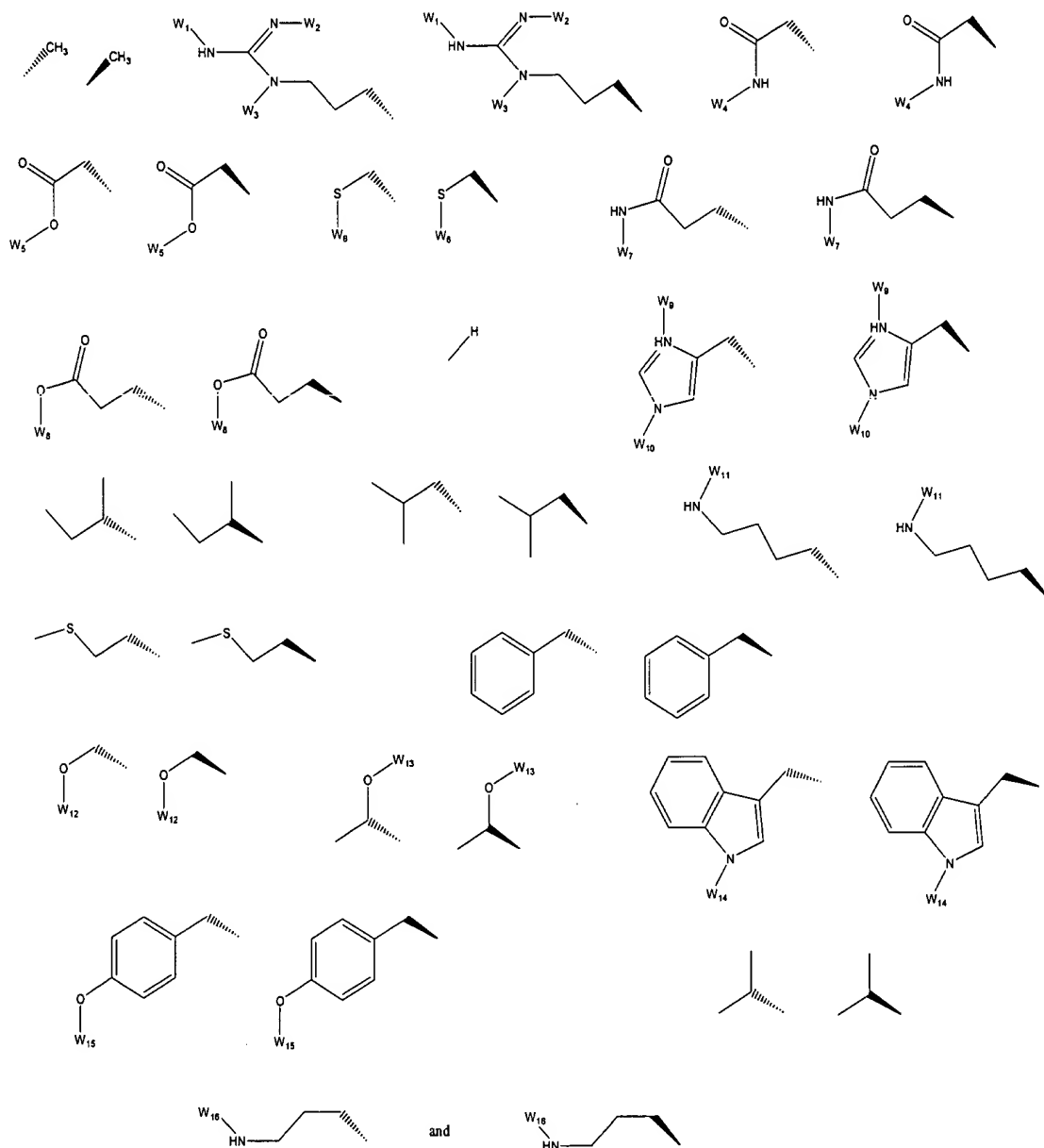


wherein

X_2 is $-\text{CH}-$, $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$;

when X_2 is $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$, R_2 is absent;

when X_2 is $-\text{CH}-$, R_2 is a radical independently selected from the group consisting of

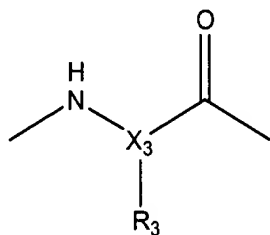


Fragment A₃ is:

(3-i) *D*-proline, *L*-proline, *D*-4-hydroxyproline, *L*-4-hydroxyproline, *D*-4-tert-butoxyproline, *L*-4-tert-butoxyproline; or

(3-ii)

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

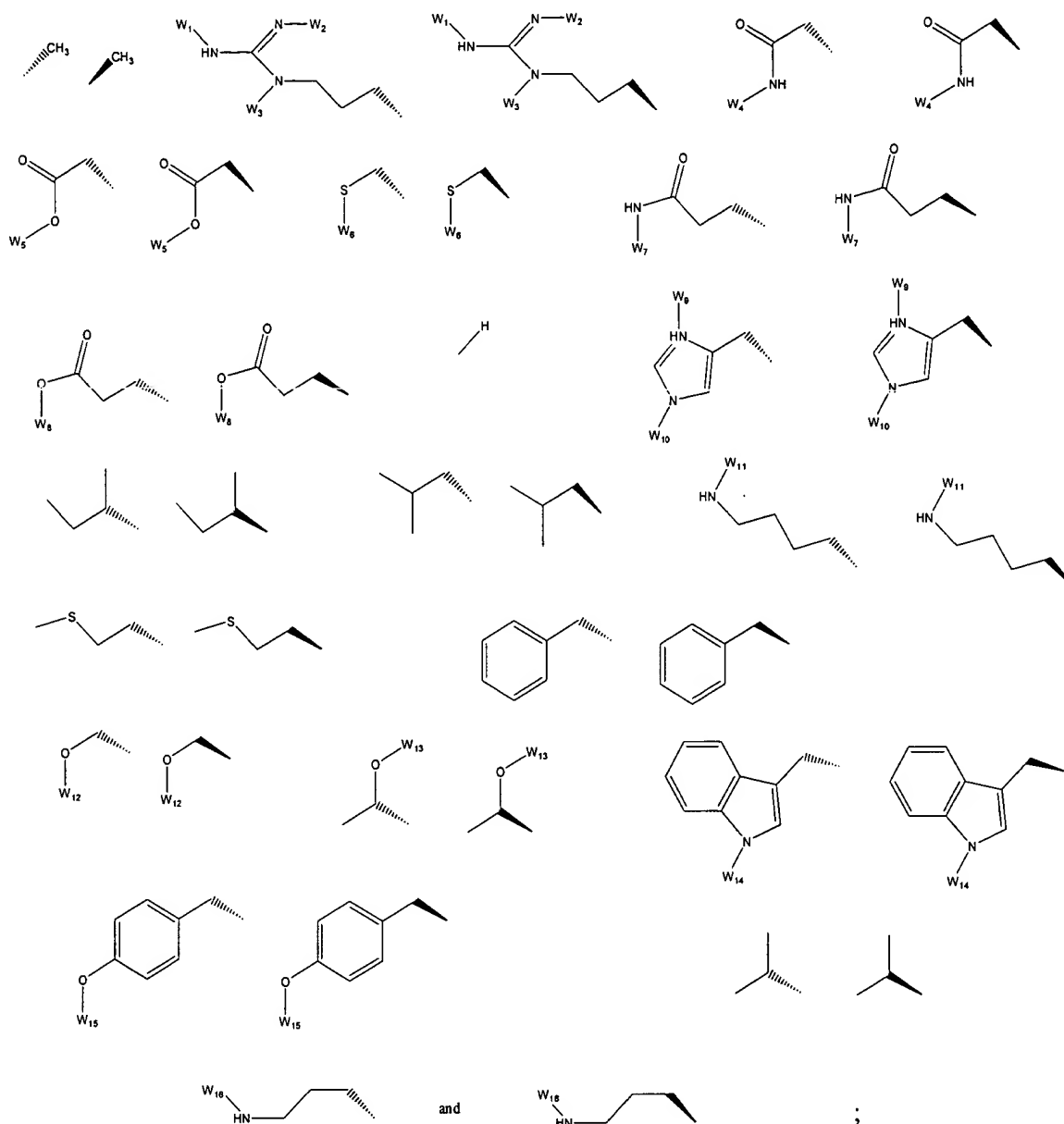


wherein

X_3 is $-\text{CH}-$, $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$;

when X_3 is $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$, R_3 is absent;

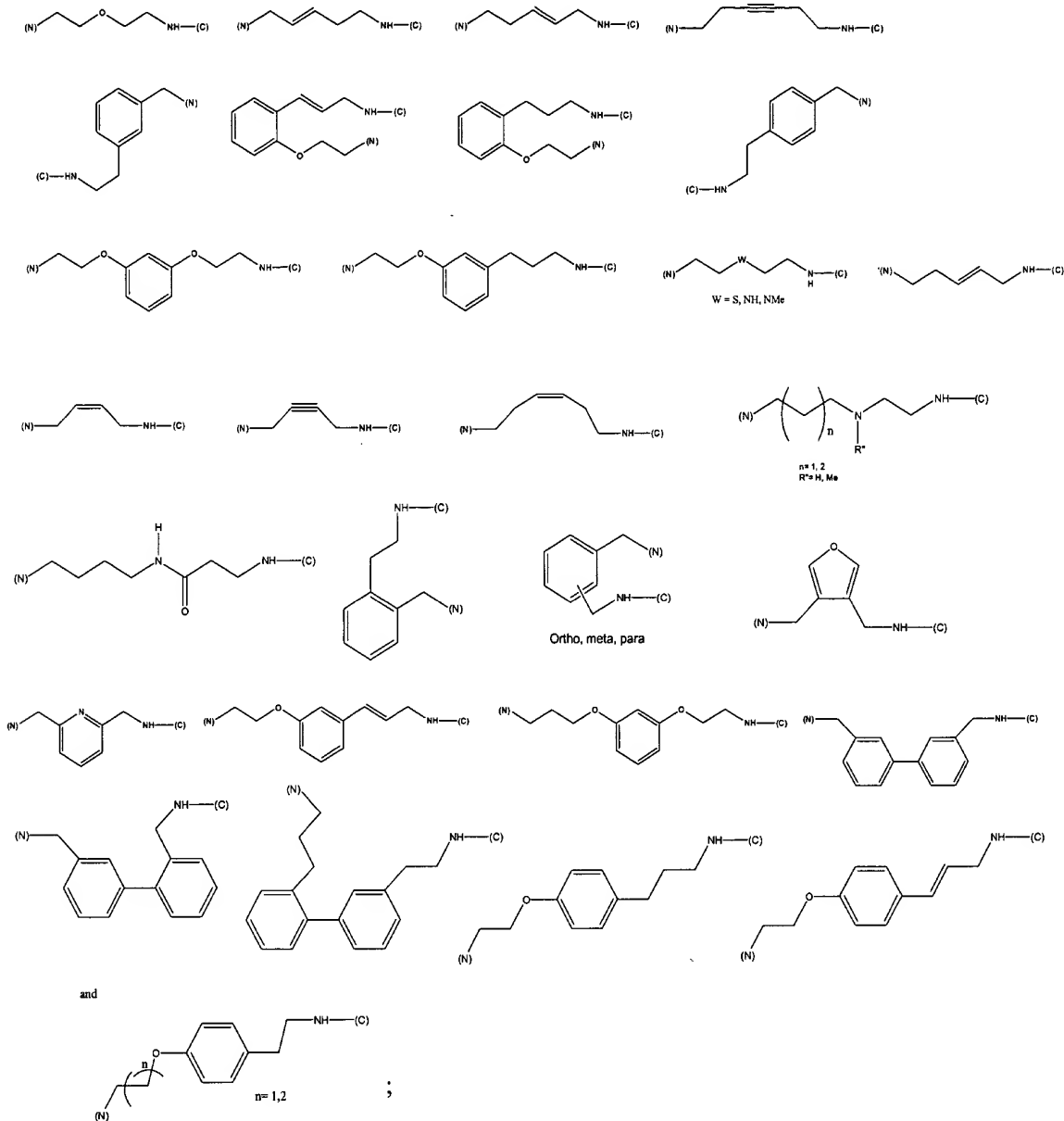
when X_3 is $-\text{CH}-$, R_3 is a radical independently selected from the group consisting of



W_1 to W_{16} are each selected from the group consisting of hydrogen and protecting groups used for orthogonal protection in peptide synthesis;

Fragment T is a radical selected from the group consisting of:

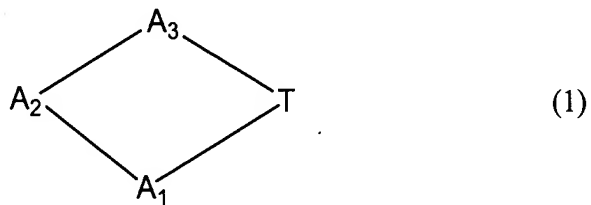
{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



wherein (N) indicates the site of a covalent bond to the nitrogen atom of A₁ of formula (1) and (C) indicates the site of a covalent bond to the carbonyl carbon of A₃ of formula (1).

Claim 36 (new): A macrocyclic compound of the formula (1):

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

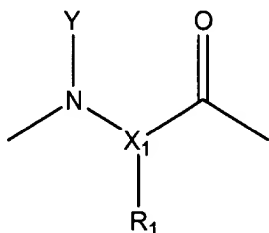


and it's pharmaceutically acceptable salts,

wherein

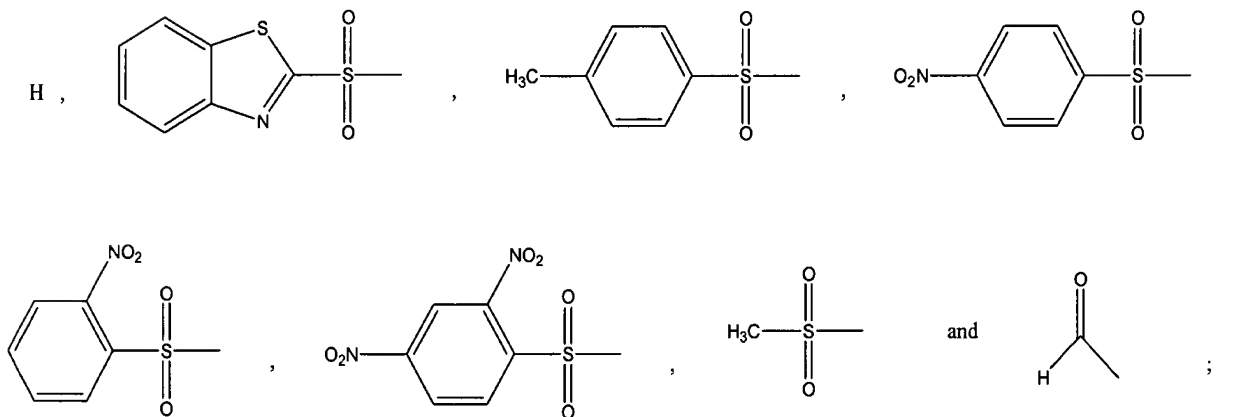
Fragment A₁ is:

(1-i)



wherein

Y is selected from the group consisting of

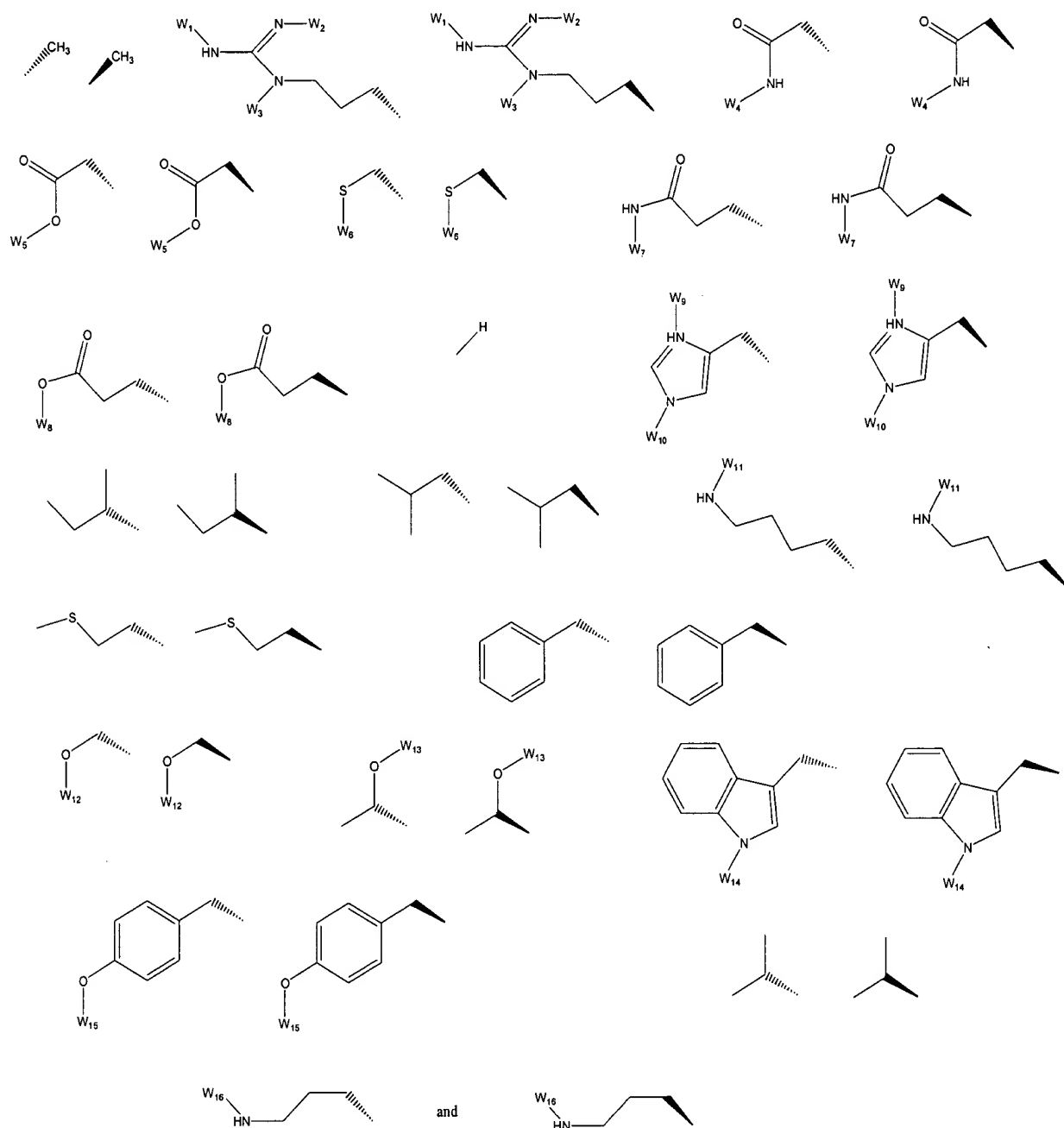


X₁ is -CH-, -(CH₂)₂- or -(CH₂)₃-;

when X₁ is -(CH₂)₂- or -(CH₂)₃-, R₁ is absent;

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

when X_1 is $-\text{CH}-$, R_1 is a radical independently selected from the group consisting of

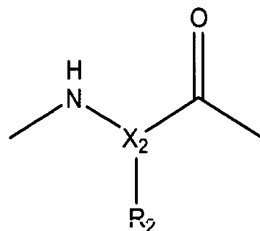


Fragment A_2 is:

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

(2-i) *D*-proline, *L*-proline, *D*-4-hydroxyproline, *L*-4-hydroxyproline, *D*-4-tert-butoxyproline, *L*-4-tert-butoxyproline; or

(2-ii)

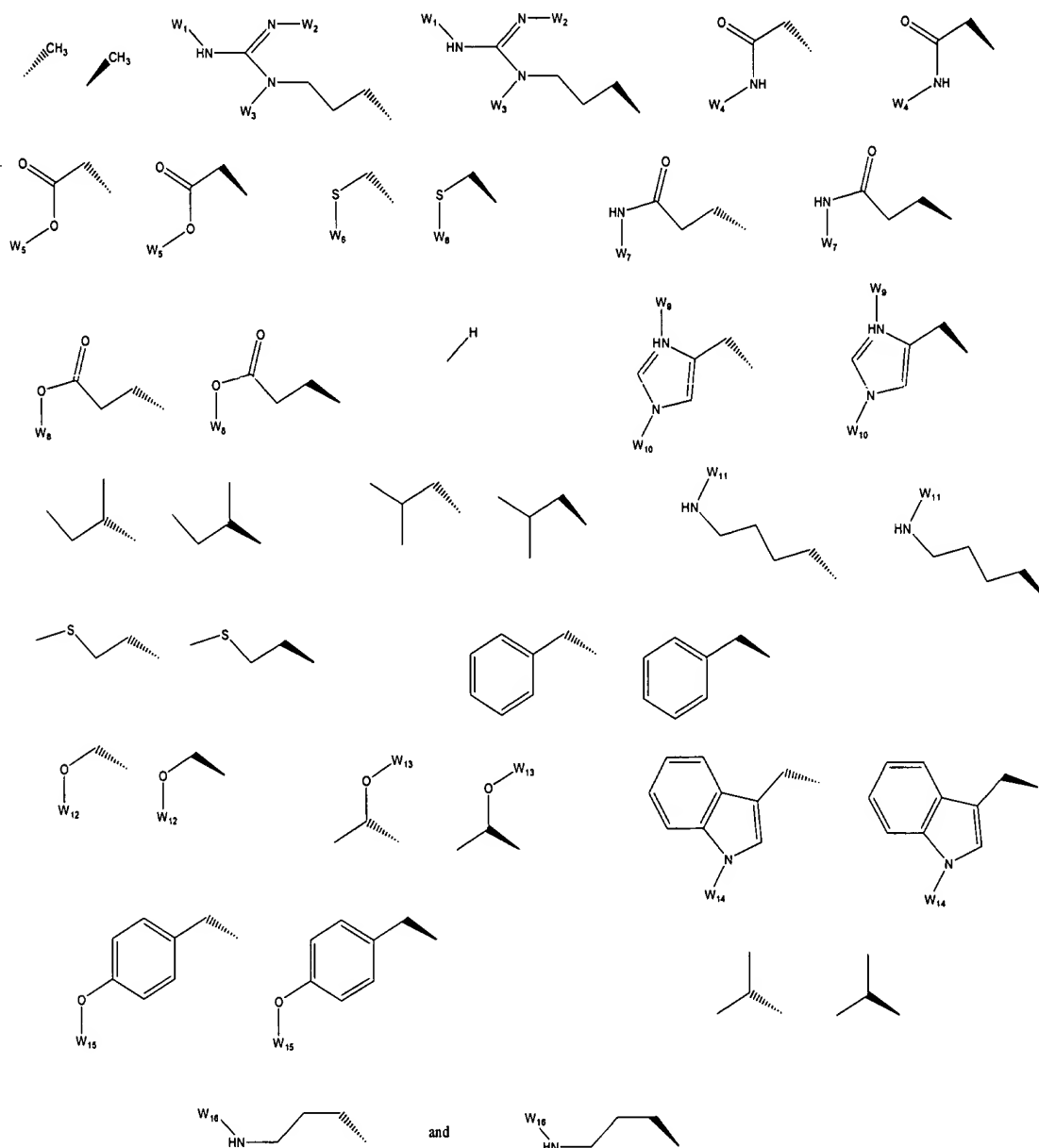


wherein

X₂ is -CH-, -(CH₂)₂- or -(CH₂)₃-;

when X₂ is -(CH₂)₂- or -(CH₂)₃-, R₂ is absent;

when X₂ is -CH-, R₂ is a radical independently selected from the group consisting of

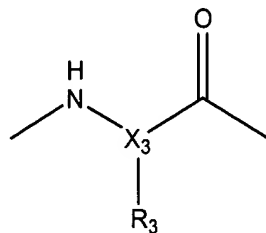


Fragment A₃ is:

(3-i) *D*-proline, *L*-proline, *D*-4-hydroxyproline, *L*-4-hydroxyproline, *D*-4-tert-butoxyproline, *L*-4-tert-butoxyproline; or

(3-ii)

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

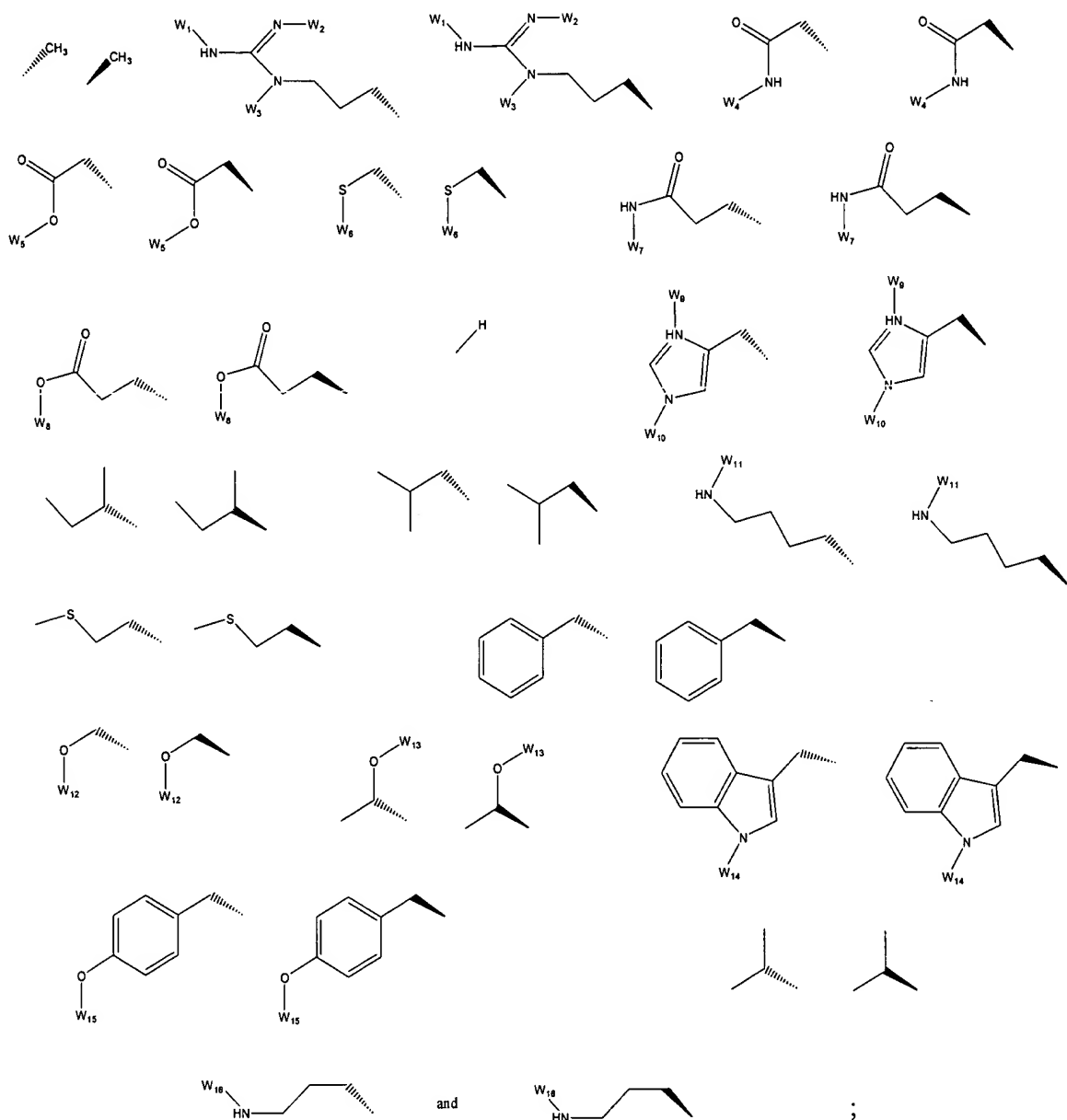


wherein

X_3 is $-\text{CH}-$, $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$;

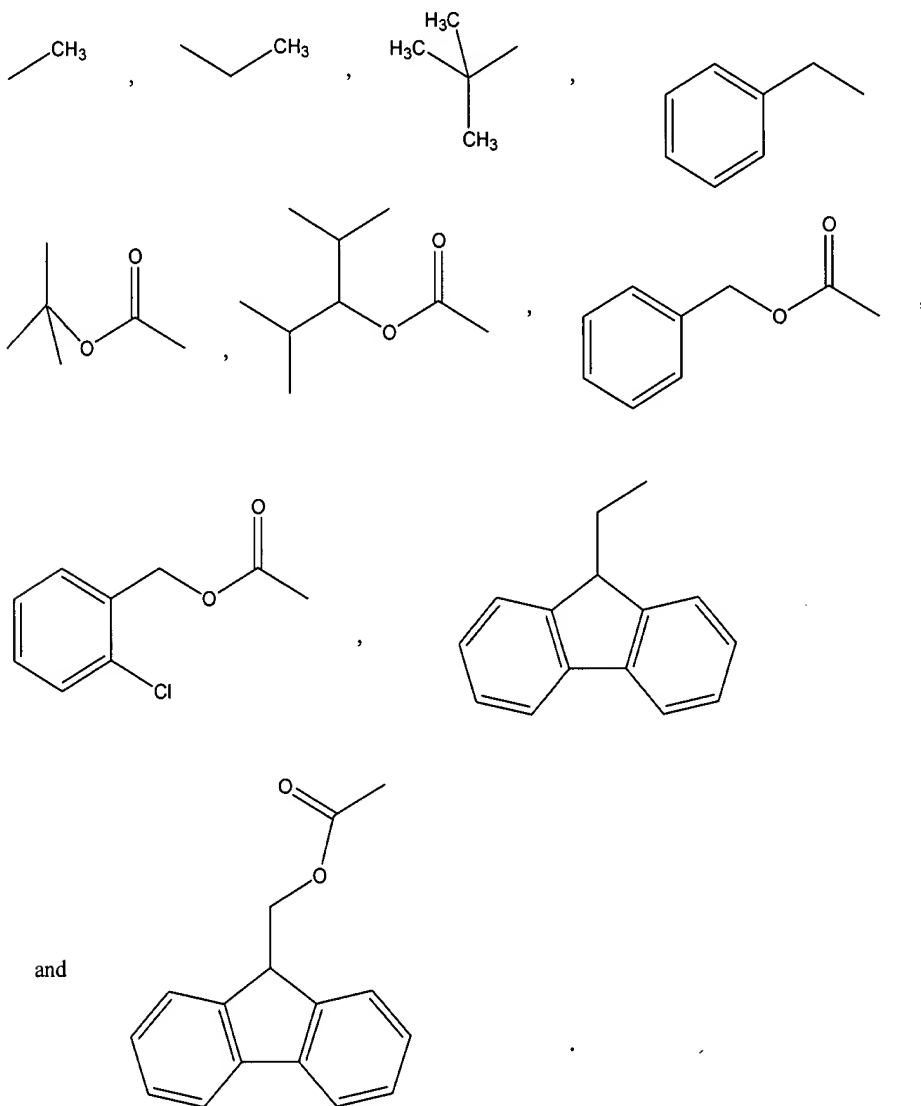
when X_3 is $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$, R_3 is absent;

when X_3 is $-\text{CH}-$, R_3 is a radical independently selected from the group consisting of



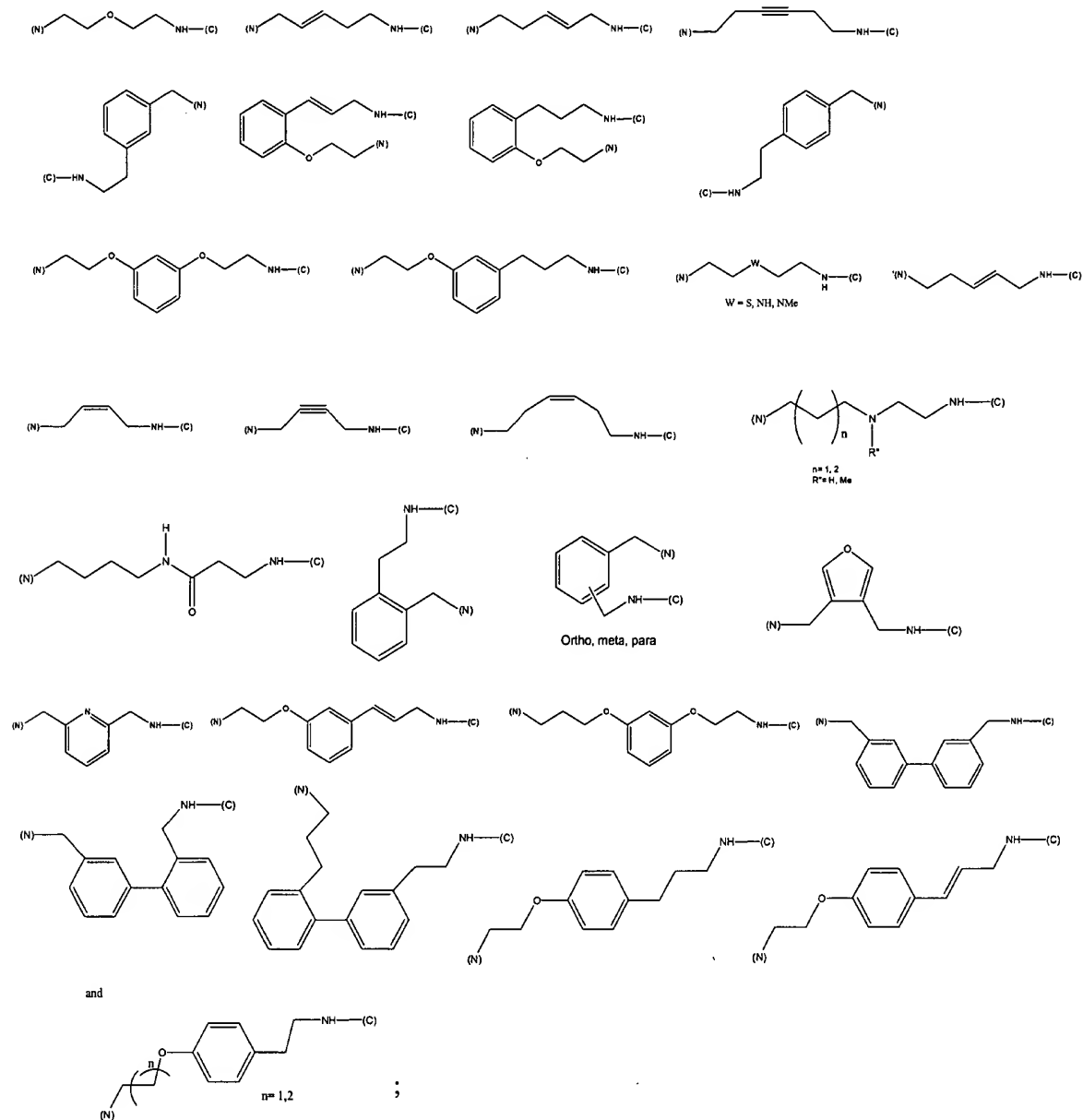
W_1 to W_{16} are each selected from the group consisting of hydrogen and a compatible protecting group chosen from:

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



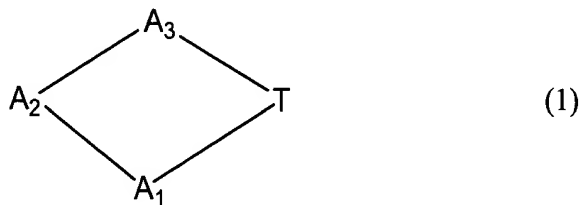
Fragment T is a radical selected from the group consisting of:

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



Claim 37 (new): A macrocyclic compound of the formula (1):

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

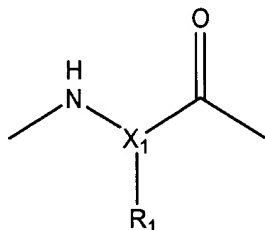


and it's pharmaceutically acceptable salts,

wherein

Fragment A₁ is:

(1-i)



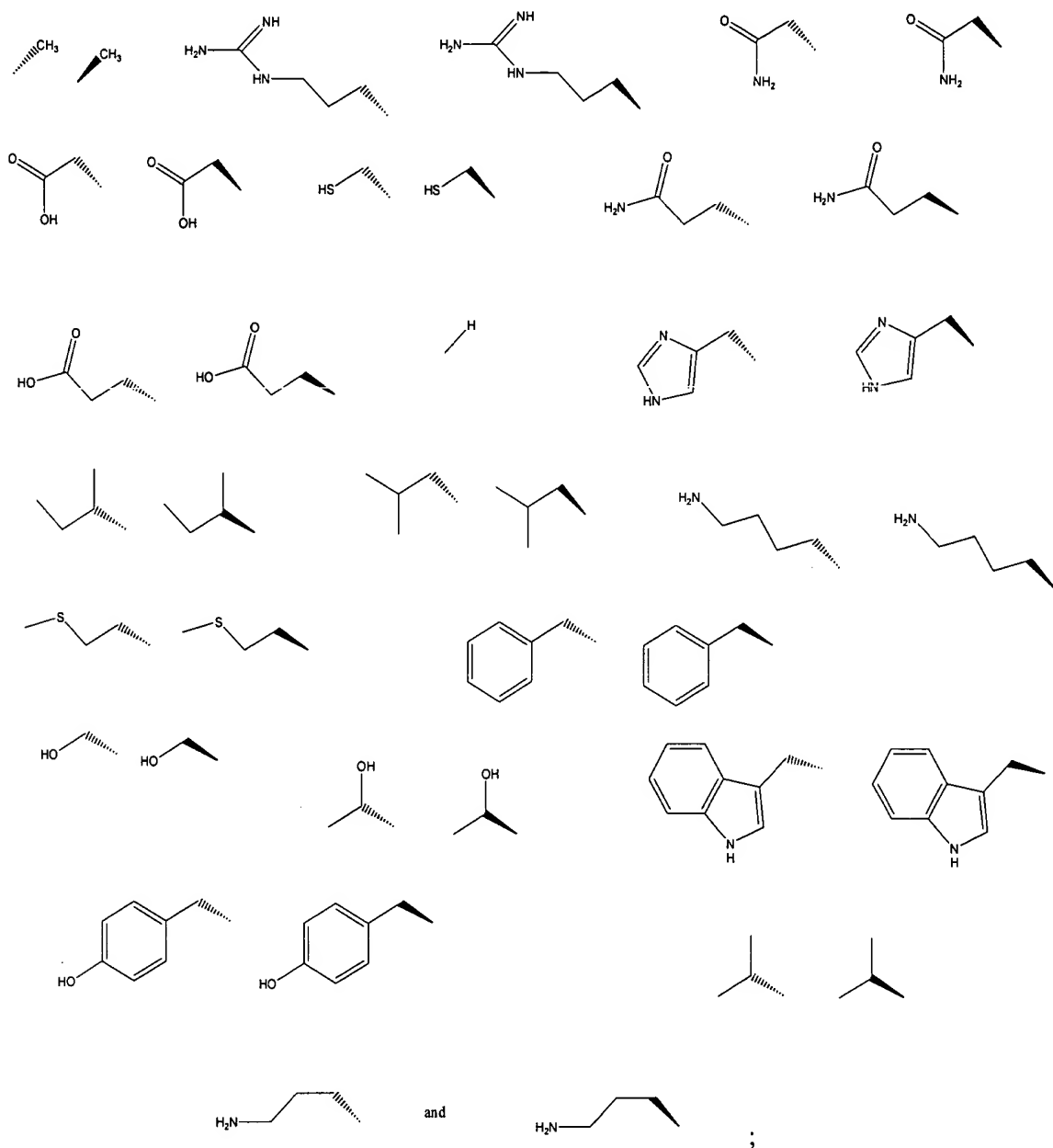
wherein

X₁ is -CH-, -(CH₂)₂- or -(CH₂)₃-;

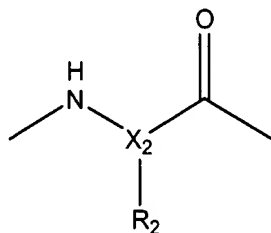
when X₁ is -(CH₂)₂- or -(CH₂)₃-, R₁ is absent;

when X₁ is -CH-, R₁ is a radical independently selected from the group consisting of:

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



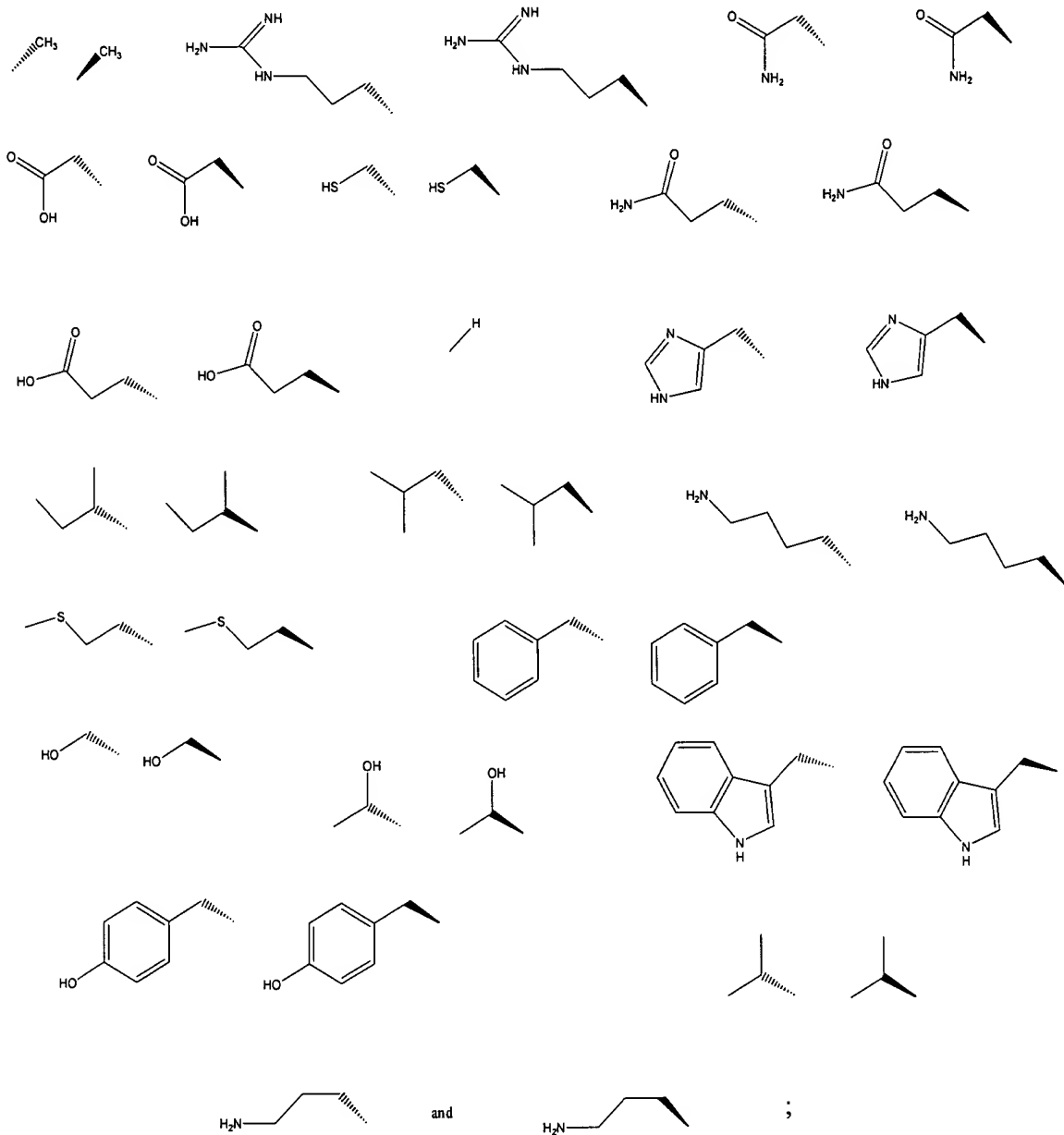
wherein

X_2 is $-\text{CH}-$, $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$;

when X_2 is $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$, R_2 is absent;

when X_2 is $-\text{CH}-$, R_2 is a radical independently selected from the group consisting of

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }

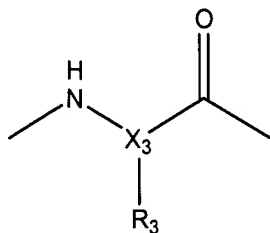


Fragment A₃ is:

(3-i) *D*-proline, *L*-proline, *D*-4-hydroxyproline, *L*-4-hydroxyproline; or

(3-ii)

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }



wherein

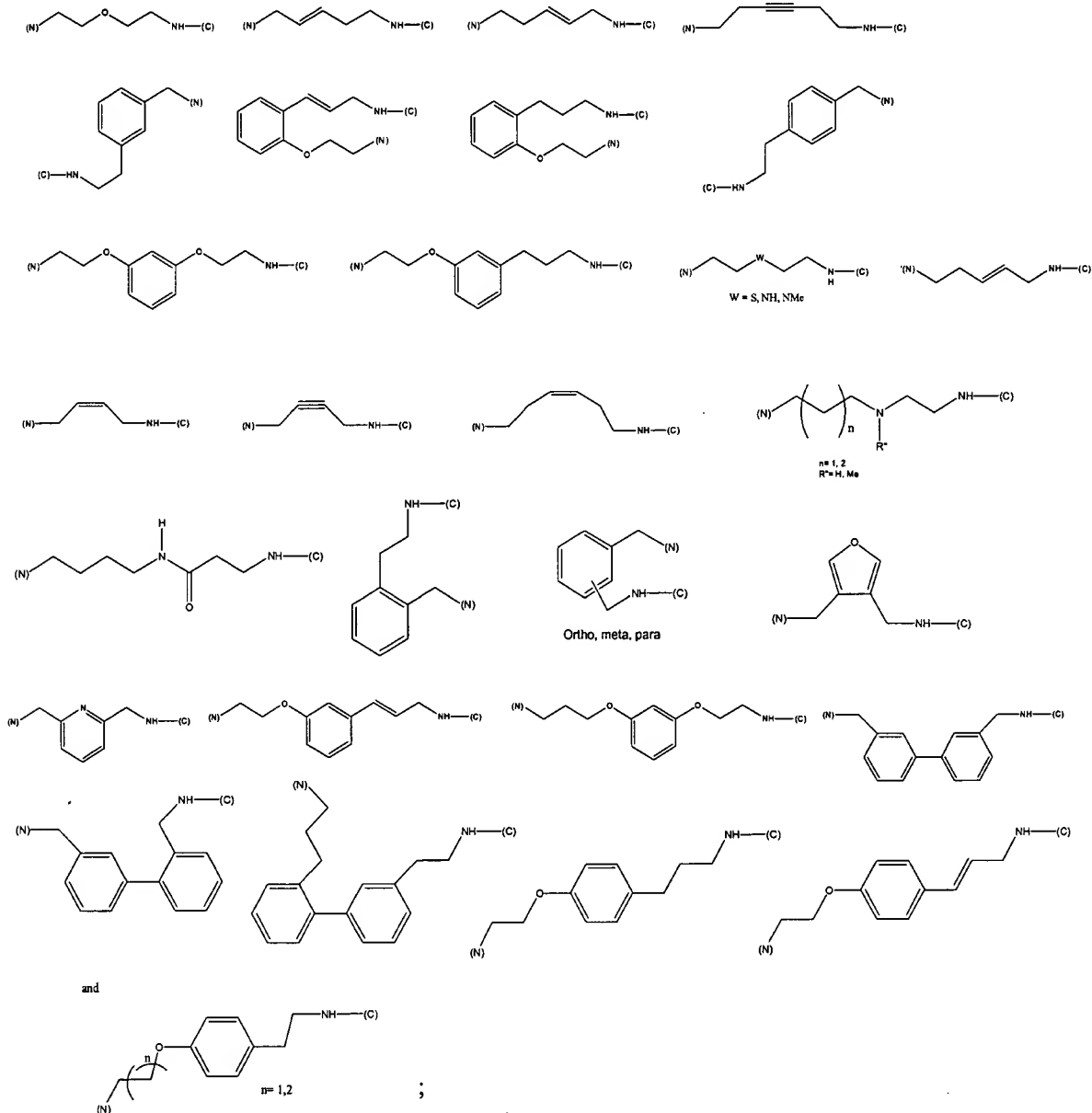
X_3 is $-\text{CH}-$, $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$;

when X_3 is $-(\text{CH}_2)_2-$ or $-(\text{CH}_2)_3-$, R_3 is absent;

when X_3 is $-\text{CH}-$, R_3 is a radical independently selected from the group consisting of



Page 24



wherein (N) indicates the site of a covalent bond to the nitrogen atom of A₁ of formula (1) and (C) indicates the site of a covalent bond to the carbonyl carbon of A₃ of formula (1).

{W:\06670\000H748-000\00080988.DOC *06670000H748-000* }